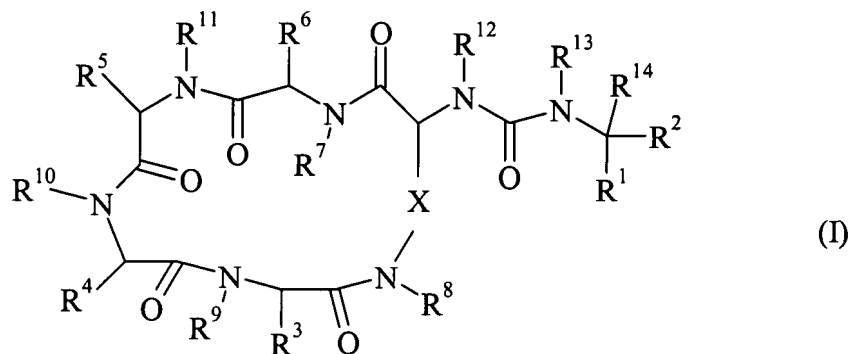


IN THE CLAIMS:

Claim 1 (**currently amended**): A method for the treatment or prophylaxis of a disease or medical condition wherein inhibition of carboxypepsidase U is beneficial, said method comprising administering to a warm-blooded animal in need thereof an effective amount~~The use~~ of a compound of formula (I):



wherein:

X is (CH₂)_mY(CH₂)_n;

m and n are, independently, 1, 2, 3, 4, 5 or 6; provided that m + n is not more than 6;

Y is a bond, O, S(O)_p, or S-S;

R¹ is CO₂R¹⁵ or a carboxylic acid isostere such as S(O)₂OH, S(O)₂NHR¹⁵, PO(OR¹⁵)OH, PO(OR¹⁵)NH₂, B(OR¹⁵)₂, PO(R¹⁵)OH, PO(R¹⁵)NH₂ or tetrazole;

R², R³, R⁴, R⁵ and R⁶ are, independently, hydrogen, C₁₋₆ alkyl (optionally substituted by halogen, hydroxy, cyano, SH, S(O)₃H, S(O)_q(C₁₋₆ alkyl), OC(O)(C₁₋₄ alkyl), CF₃, C₁₋₄ alkoxy, OCF₃, COOH, CONH₂, CONH(C₁₋₆ alkyl), NH₂, CNH(NH₂), or NHCNH(NH₂)), C₃₋₆ cycloalkyl(C₁₋₄)alkyl (wherein the cycloalkyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)), heterocyclyl(C₁₋₄)alkyl (wherein the heterocyclyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)), phenyl(C₁₋₄)alkyl (wherein the phenyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)) or heteroaryl(C₁₋₄)alkyl (wherein the heteroaryl ring is

optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂));

p and q are, independently, 0, 1 or 2;

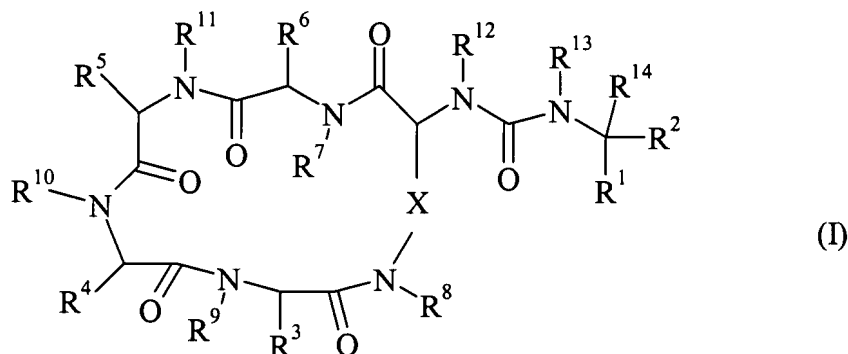
R⁷, R⁸, R⁹, R¹⁰, R¹¹, R¹² and R¹³ are, independently, H or C₁₋₄ alkyl;

R¹⁴ is H or C₁₋₄ alkyl; and,

R¹⁵ is H or C₁₋₄ alkyl;

or a pharmaceutically acceptable salt thereof, ~~or solvate thereof, or a solvate of such a salt; in a method of manufacturing a medicament for the treatment or prophylaxis of a condition wherein inhibition of carboxypeptidase U is beneficial.~~

Claim 2 (currently amended): A compound of formula (I):



wherein:

X is (CH₂)₄;

R¹ is CO₂R¹⁵;

R² is C₁₋₆ alkyl, benzyl, straight-chain C₁₋₆ alkyl substituted at its terminus by NH₂, CNH(NH₂)₁₋₂ or NHCNH(NH₂) or (6-aminopyridin-3-yl)methyl; C₃₋₆ cycloalkyl substituted by NH₂, CNH(NH₂) or NHCNH(NH₂); heterocyclyl containing at least one nitrogen atom; non-nitrogen containing heterocyclyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); heteroaryl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); phenyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); heteroaryl(C₁₋₄)alkyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); phenyl(C₁₋₄)alkyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); or C₃₋₆ cycloalkyl(C₁₋₄)alkyl substituted with NH₂,

CNH(NH₂) or NHCNH(NH₂); all of the above rings being optionally further substituted by one or more of: halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy or OCF₃; one of R³, R⁴, R⁵ and R⁶ is independently, hydrogen, heteroaryl(C₁₋₄)alkyl (wherein the heteroaryl ring is optionally substituted by one or more of halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)); and the others are, independently, hydrogen, C₁₋₆ alkyl (optionally substituted by halogen, hydroxy, cyano, SH, S(O)₃H, S(O)_q(C₁₋₆ alkyl), OC(O)(C₁₋₄ alkyl), CF₃, C₁₋₄ alkoxy, OCF₃, COOH, CONH₂, CONH(C₁₋₆ alkyl), NH₂, CNH(NH₂), or NHCNH(NH₂)), C₃₋₆ cycloalkyl(C₁₋₄)alkyl (wherein the cycloalkyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)), heterocyclyl(C₁₋₄)alkyl (wherein the heterocyclyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)), phenyl(C₁₋₄)alkyl (wherein the phenyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)) or heteroaryl(C₁₋₄)alkyl (wherein the heteroaryl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)); p and q are, independently, 0, 1 or 2; R⁷, R⁸, R⁹, R¹⁰, R¹¹, R¹² and R¹³ are, independently, H or C₁₋₄ alkyl; R¹⁴ is H or C₁₋₄ alkyl; and, R¹⁵ is H or C₁₋₄ alkyl;

or a pharmaceutically acceptable salt ~~thereof or solvate thereof, or a solvate of such a salt.~~

Claim 3 (**currently amended**): ~~A~~ The compound of formula (I) ~~or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt.~~ as claimed in claim 2 wherein:

X is (CH₂)₄;

R¹ is CO₂R¹⁵;

R² is straight-chain C₁₋₆ alkyl substituted at its terminus by NH₂, CNH(NH₂) or NHCNH(NH₂); C₃₋₆ cycloalkyl substituted by NH₂, CNH(NH₂) or NHCNH(NH₂); heterocyclyl containing at least one nitrogen atom; non-nitrogen containing heterocyclyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); heteroaryl substituted with NH₂,

CNH(NH₂) or NHCNH(NH₂); phenyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); heteroaryl(C₁₋₄)alkyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); phenyl(C₁₋₄)alkyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); or C₃₋₆ cycloalkyl(C₁₋₄)alkyl substituted with NH₂, CNH(NH₂) or NHCNH(NH₂); all of the above rings being optionally further substituted by one or more of: halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy or OCF₃;

one of R³, R⁴, R⁵ and R⁶ is independently, hydrogen, heteroaryl(C₁₋₄)alkyl (wherein the heteroaryl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)); and the others are, independently, hydrogen, C₁₋₆ alkyl (optionally substituted by halogen, hydroxy, cyano, SH, S(O)₃H, S(O)₄(C₁₋₆ alkyl), OC(O)(C₁₋₄ alkyl), CF₃, C₁₋₄ alkoxy, OCF₃, COOH, CONH₂, CONH(C₁₋₆ alkyl), NH₂, CNH(NH₂), or NHCNH(NH₂)), C₃₋₆ cycloalkyl(C₁₋₄)alkyl (wherein the cycloalkyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)), heterocyclyl(C₁₋₄)alkyl (wherein the heterocyclyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)), phenyl(C₁₋₄)alkyl (wherein the phenyl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂)) or heteroaryl(C₁₋₄)alkyl (wherein the heteroaryl ring is optionally substituted by halogen, hydroxy, cyano, C₁₋₄ alkyl, CF₃, C₁₋₄ alkoxy, OCF₃, NH₂, CNH(NH₂) or NHCNH(NH₂));

p and q are, independently, 0, 1 or 2;

R⁷, R⁸, R⁹, R¹⁰, R¹¹, R¹² and R¹³ are, independently, H or C₁₋₄ alkyl;

R¹⁴ is H or C₁₋₄ alkyl; and,

R¹⁵ is H or C₁₋₄ alkyl;

or a pharmaceutically acceptable salt thereof or solvate thereof, or a solvate of such a salt.

Claim 4 (currently amended): A ~~The~~ compound of formula (I) ~~or a pharmaceutically acceptable salt or solvate thereof, or a solvate of such a salt~~ as claimed in claim 2 ~~or 3~~ wherein:

R¹ is CO₂R¹⁵;

R^2 is straight-chain C_{1-6} alkyl substituted at its terminus by NH_2 , $CNH(NH_2)$ or $NHCNH(NH_2)$; C_4 alkyl (such as $CH(CH_3)CH_2CH_3$ or $CH_2CH(CH_3)_2$); or (aminopyridinyl)methyl (for example (6-aminopyridin-3-yl)methyl); one of R^3 and R^4 is (indol-3-yl) CH_2 optionally substituted by halo or hydroxy; and the other is benzyl (optionally substituted by halo or hydroxy) or C_4 alkyl (such as $CH(CH_3)CH_2CH_3$ or $CH_2CH(CH_3)_2$); or R^3 and R^4 are both methyl; R^5 and R^6 are, independently, C_{1-6} alkyl (for example CH_3 , $CH(CH_3)_2$, $CH(CH_3)CH_2CH_3$ or $CH_2CH(CH_3)_2$); R^7 , R^8 , R^9 , R^{11} , R^{12} , R^{13} and R^{14} are H; R^{10} is C_{1-4} alkyl; and, R^{15} is H or C_{1-4} alkyl;

or a pharmaceutically acceptable salt thereof.

Claim 5 (**currently amended**): The method of claim 1 ~~A compound as claimed in any one of claims 2 to 4~~ wherein X is $(CH_2)_4$.

Claim 6 (**currently amended**): The method of claim 1 ~~A compound as claimed in any one of claims 2 to 5~~ wherein R^1 is CO_2R^{15} in which R^{15} is H or C_{1-4} alkyl.

Claim 7 (**currently amended**): A ~~The compound as claimed in claim 2~~ ~~any one of claims 2 to 6~~ wherein R^2 is straight-chain C_{1-6} alkyl substituted at its terminus by NH_2 , $CNH(NH_2)$ or $NHCNH(NH_2)$; C_4 alkyl (such as $CH(CH_3)CH_2CH_3$ or $CH_2CH(CH_3)_2$); or (aminopyridinyl)methyl.

Claim 8 (**currently amended**): A ~~The compound as claimed in claim 2~~ ~~any one of claims 2 to 4~~ wherein R^2 is C_{1-6} alkyl ($CH(CH_3)CH_2CH_3$ or $CH_2CH(CH_3)_2$), benzyl, or straight-chain C_{1-6} alkyl substituted at its terminus by NH_2 , $CNH(NH_2)$, $NHCNH(NH_2)$ or (6-aminopyridin-3-yl)methyl.

Claim 9 (**currently amended**): A ~~The~~ compound as claimed in claim 2 ~~any one of claims 2 to 8~~ wherein R^2 is straight-chain C_{1-6} alkyl substituted at its terminus by NH_2 , $CNH(NH_2)$, $NHCNH(NH_2)$ or (6-aminopyridin-3-yl)methyl.

Claim 10 (**currently amended**): A ~~The~~ compound as claimed in claim 2 ~~any one of claims 2 to 10~~ wherein R^3 is $CH_2indolyl_1$ -(wherein the indolyl is optionally substituted by one or more of: halogen or hydroxy, C_{1-4} alkyl or benzyl (optionally substituted by halogen or hydroxy)).

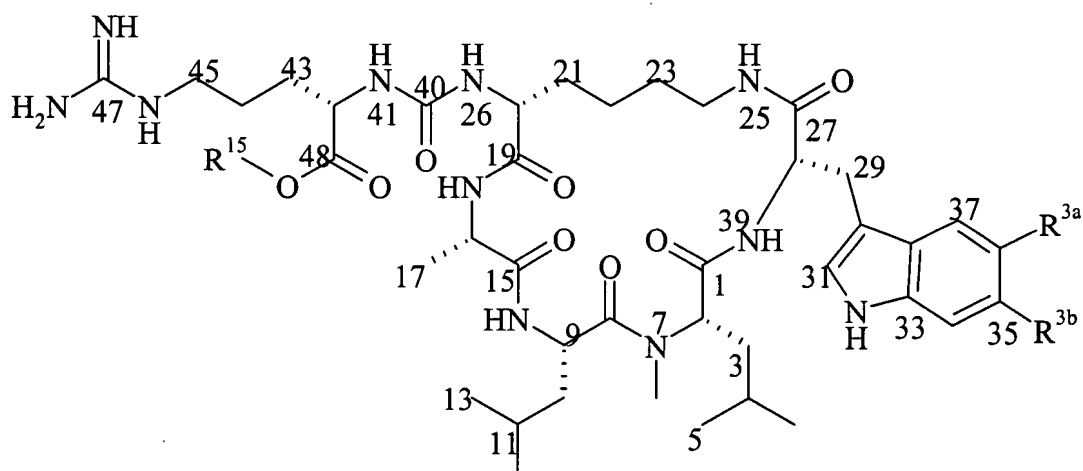
Claim 11 (**currently amended**): A ~~The~~ compound as claimed in claim 2 ~~any one of claims 2 to 10~~ wherein R^4 is $CH_2indolyl_1$ -(wherein the indolyl is optionally substituted by one or more of: halogen or hydroxy, C_{1-6} alkyl ($CH(CH_3)CH_2CH_3$ or $CH_2CH(CH_3)_2$) or benzyl (optionally substituted by halogen or hydroxy)).

Claim 12 (**currently amended**): A ~~The~~ compound as claimed in claim 2 ~~any one of claims 2 to 11~~ wherein R^5 and R^6 are, independently, C_{1-6} alkyl (~~such as methyl, iso-propyl, $CH(CH_3)CH_2CH_3$ or $CH_2CH(CH_3)_2$.~~

Claim 13 (**currently amended**): A ~~The~~ compound as claimed in claim 2 ~~any one of claims 2 to 12~~ wherein R^7 , R^8 , R^9 , R^{11} , R^{12} , R^{13} and R^{14} are all H.

Claim 14 (**currently amended**): A ~~The~~ compound as claimed in claim 2 ~~any one of claims 2 to 4~~ wherein R^{10} is C_{1-4} alkyl.

Claim 15 (**currently amended**): A ~~The~~ compound as claimed in claim 2 which is a compound of the following formula



in which

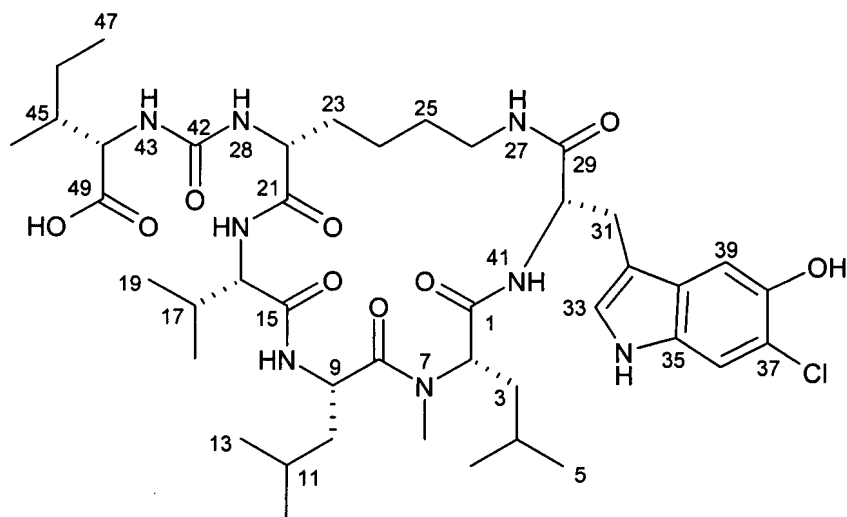
R^{3a} is H, R^{3b} is H and R¹⁵ is H;

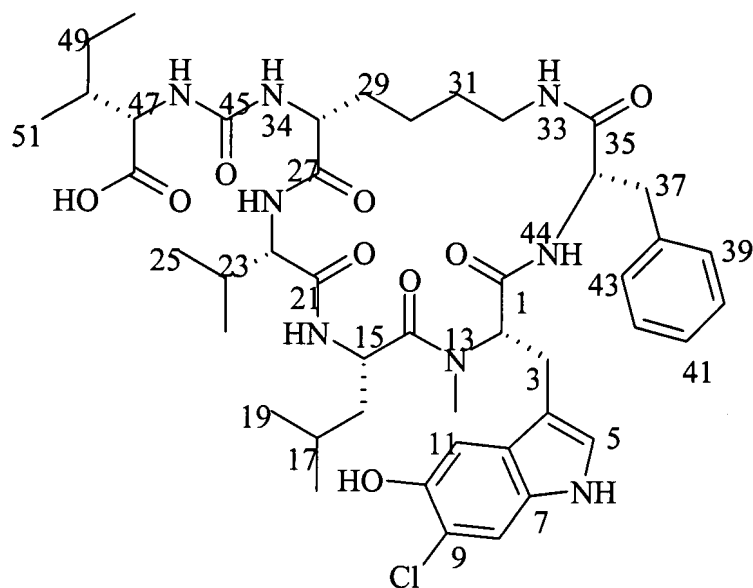
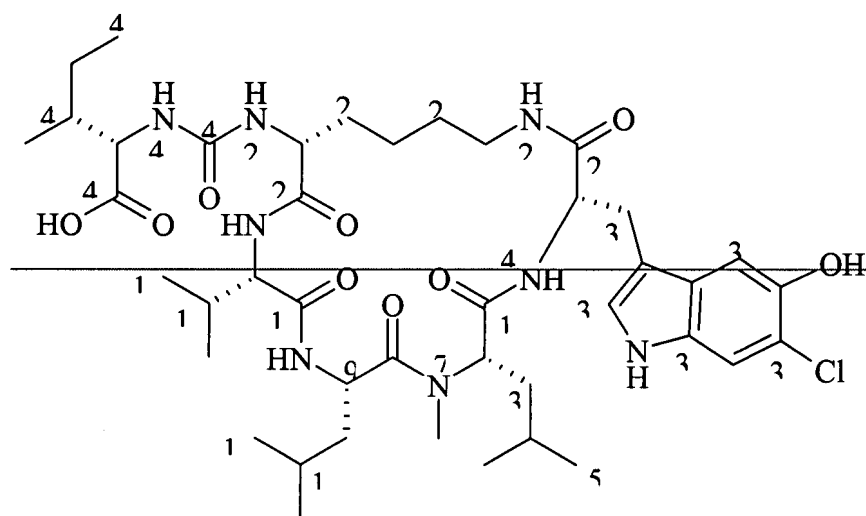
R^{3a} is OH, R^{3b} is Cl and R¹⁵ is H;

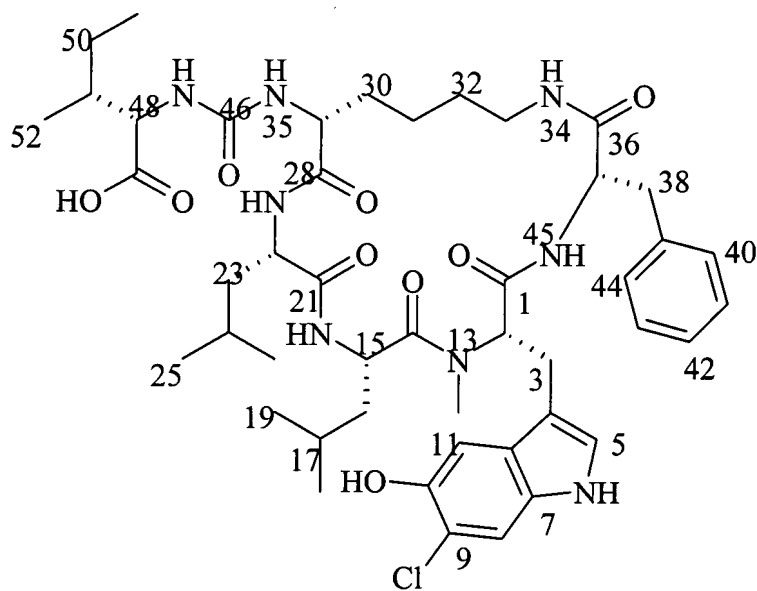
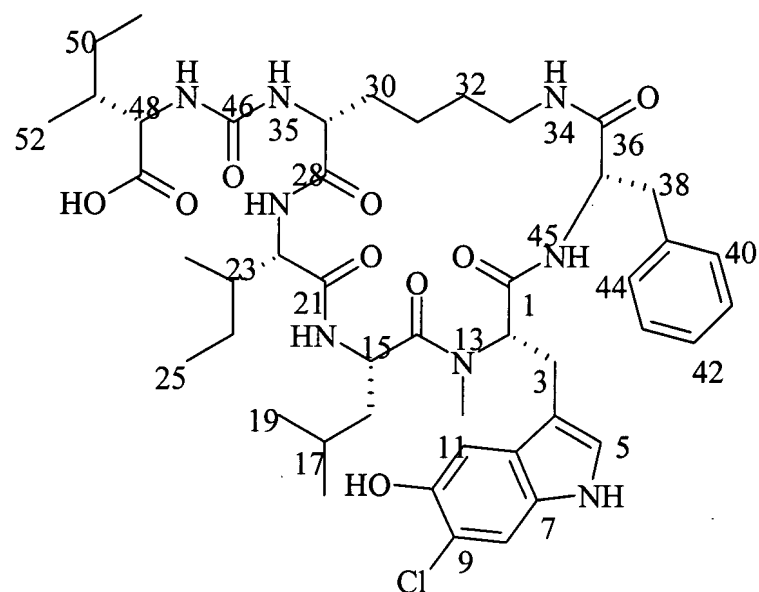
R^{3a} is OH, R^{3b} is Cl and R^{15-is} is CH₃;

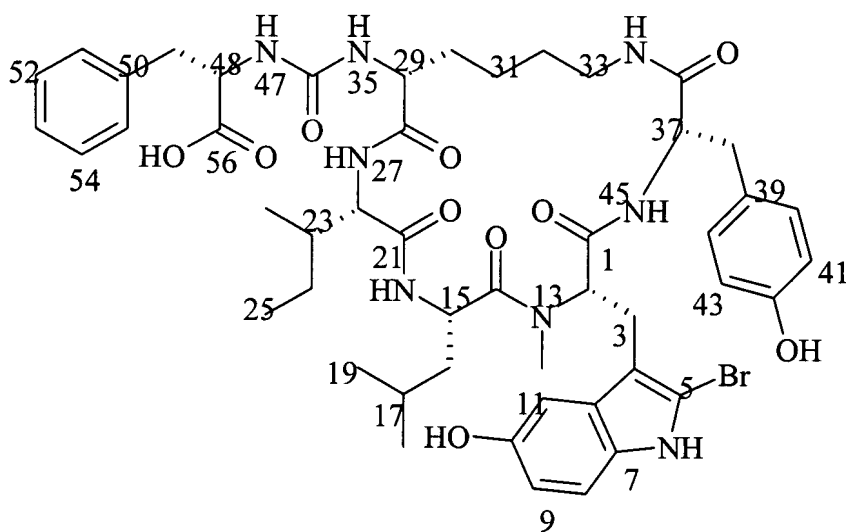
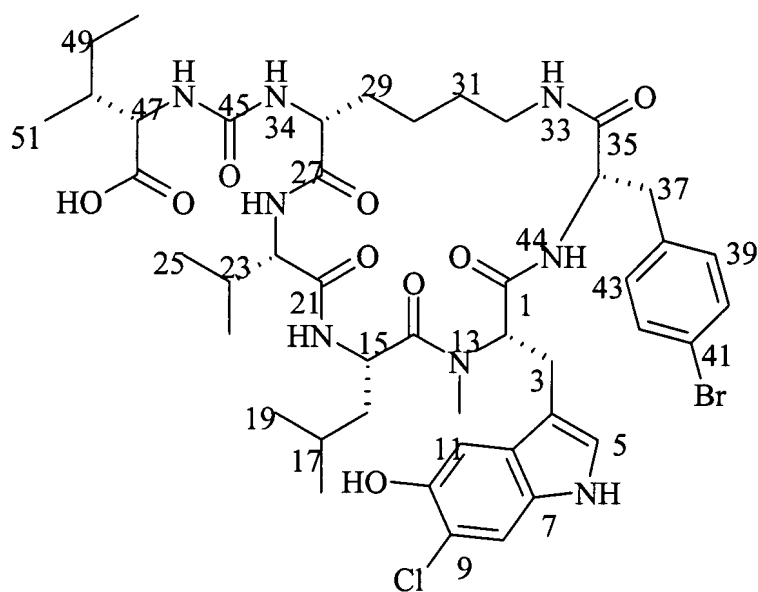
R^{3a} is H, R^{3b} is H and R^{15-is} is CH₃;

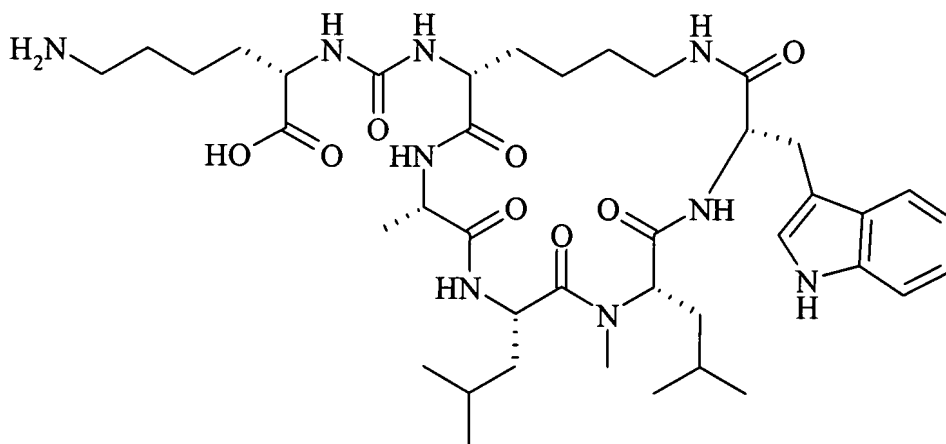
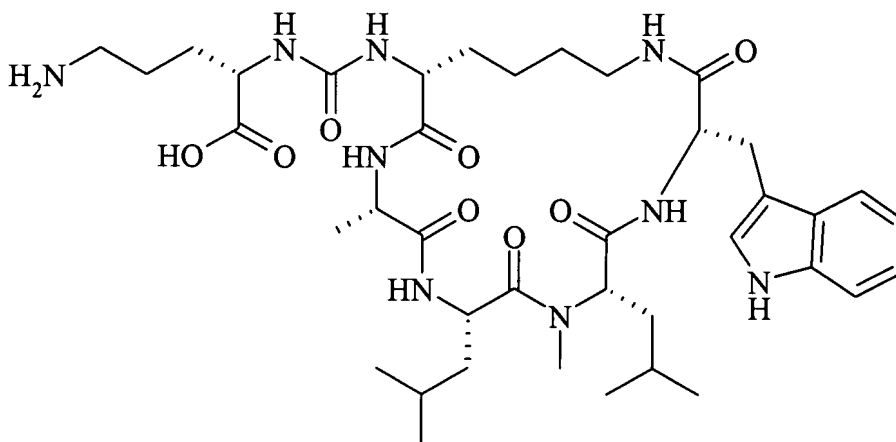
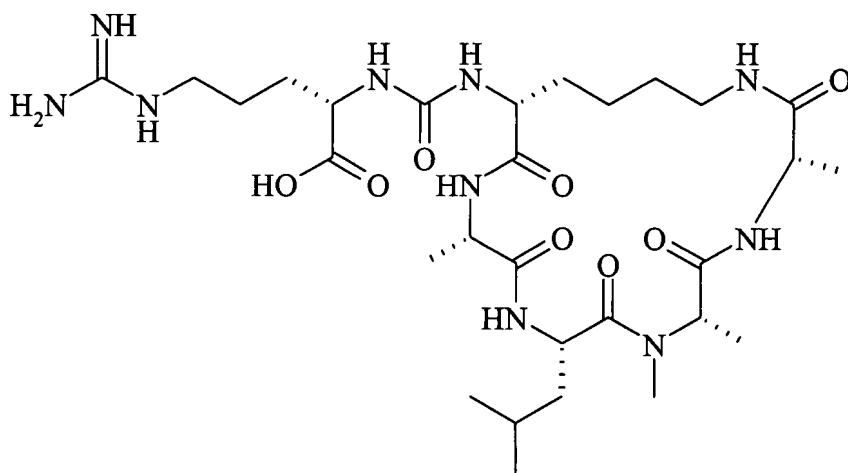
R^{3a} is H, R^{3b} is Cl and R¹⁵ is H;

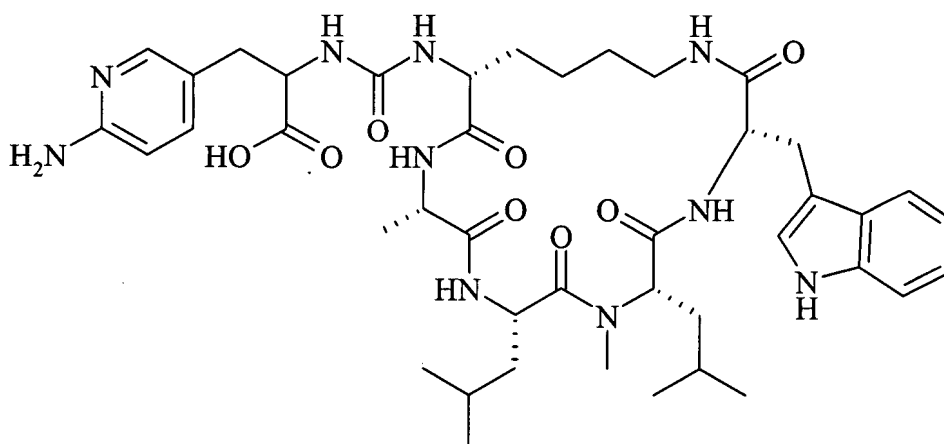




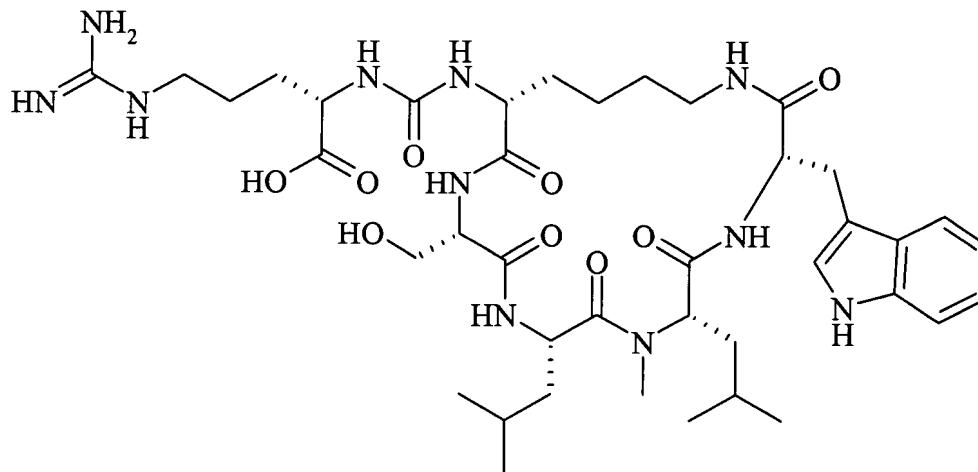








or



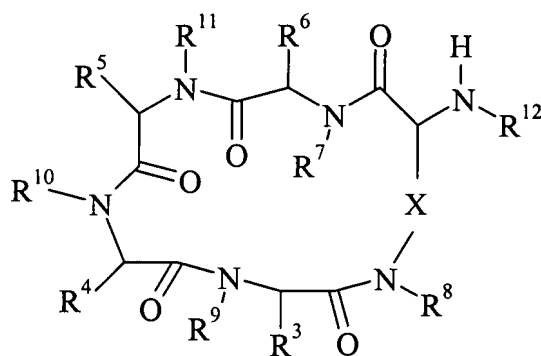
or a pharmaceutically acceptable salt thereof or solvate thereof, or a solvate of a pharmaceutically acceptable salt thereof.

Claim 16 (**currently amended**): A method for the treatment or prophylaxis of a disease or medical condition wherein inhibition of carboxypepsidase U is beneficial, said method comprising administering to a warm-blooded animal in need thereof an effective amount ~~The use of a compound of formula (I) or a pharmaceutically acceptable salt thereof or solvate thereof, or a solvate of such a salt; as claimed in claim 2 any one of claims 2 to 15 in a method of manufacturing a medicament for the treatment or prophylaxis of a condition wherein inhibition of carboxypeptidase U is beneficial.~~

Claim 17 (**currently amended**): The method-use as claimed in claim 16 wherein said disease or medical condition is selected from ~~for the manufacture of a medicament for the treatment or prophylaxis of~~ thrombosis and/or hypercoagulability in blood and/or tissues; atherosclerosis; fibrotic conditions; inflammatory diseases; or a condition which benefits from maintaining or enhancing bradykinin levels in the body of a mammal ~~(such as man)~~.

Claim 18 (**currently amended**): A pharmaceutical formulation comprising ~~containing~~ a compound of formula (I) or a pharmaceutically acceptable salt thereof ~~or solvate thereof, or a solvate of such a salt~~; as claimed in claim 2 ~~any one of claims 2 to 15~~ as active ingredient in combination with a pharmaceutically acceptable adjuvant, diluent or carrier.

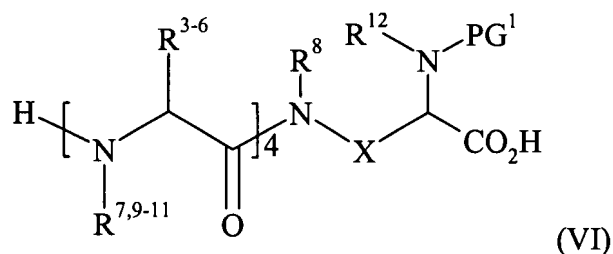
Claim 19 (**currently amended**): A compound of formula



(VII)

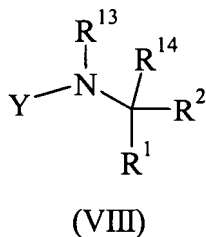
wherein R³ to R¹² and X are as defined in claim 2 ~~any one of claims 1 to 14~~.

Claim 20 (**currently amended**): A process for preparing a compound as claimed in claim 19 which comprises treating a compound of formula VI ~~in which PG1 is a suitable protecting group with a peptide coupling agent in the presence of a non-nucleophilic base in a polar aprotic solvent and then removing the protecting group.~~



in which PG¹ is a suitable protecting group with a peptide coupling agent in the presence of a non-nucleophilic base in a polar aprotic solvent and then removing the protecting group.

Claim 21 (**currently amended**): A process for preparing a compound of formula I as claimed in claim 2 ~~any one of claims 2 to 17~~ which comprises reacting a compound of formula VII as defined in claim 19 with a compound of formula VIII



in which Y is an activated ester or NY is an isocyanate group.

Claim 22 (**new**): The method as claimed in claim 1 wherein said disease or medical condition is selected from thrombosis and/or hypercoagulability in blood and/or tissues; atherosclerosis; fibrotic conditions; inflammatory diseases; or a condition which benefits from maintaining or enhancing bradykinin levels in the body of a mammal.